## **REMARKS**

Reconsideration of this application, as amended, is respectfully requested. The Applicants wish to draw the Examiner's attention to the applicants' related co-pending applications and issued patents (see Appendix A) directed to nanoparticles and methods of preparation and use thereof.

The Applicants note that the Examiner did not return an executed PTO 1449 form for the 6<sup>th</sup> Supp. IDS that was hand-delivered to the Examiner on September 9, 2002. Subsequent to the issuance of this Office action, the Applicants had filed a 7<sup>th</sup> Supp. IDS. The Applicants request that the Examiner fully execute the PTO 1449 forms for the 6<sup>th</sup> and 7<sup>th</sup> Supp. IDS and return a copy of the executed PTO 1449 forms to the undersigned representative. Copies of the initial IDS, 6<sup>th</sup> and 7<sup>th</sup> Supplemental IDS, and associated PTO 1449 forms are attached. The Examiner is requested to contact the undersigned representative if the Examiner would like to have another copy of the references.

The specification has been amended to update the priority claim. No new matter has been added to the application as a result of this amendment.

Claims 125-129 and 156 were originally pending in this application. Claims 125-129, 156, and 157 were cancelled, claims 158-161 were amended, and new claims 433-465 were added. The Examiner had indicated that claims 129 and 157 were allowed. As the Examiner will note, these claims were cancelled and reintroduced as new claims 464 and 465, respectively. The amended and new claims are fully supported in the previously pending claims, and thus do not constitute new matter. The amended and new claims are supported, for example, by the original claims and the specification at page 48, line 12 to page 50, line 2; and page 77, line 1 to page 80, line 27. Accordingly, no new matter has been introduced into the application as a result of the above amendment. Claims 158-161 and 433-465 are now pending in this application.

Turning to the office action, claims 128 was rejected under 35 U.S.C. section 112, second paragraph, for indefiniteness. Claims 125, 126 and 156 were also rejected for obviousness-type double patenting as being unpatentable over claims 31 and 33 of U.S. Patent no. 6,506,564. The aforementioned claims were cancelled and thus the aforementioned rejections are moot. Claims 125-128, 156, and 158 stand rejected under 35 U.S.C. section 102(e) as being anticipated by

Yguerabide (U.S. Patent No. 6,214,560)("Yguerabide"). The Applicants respectfully traverse this rejection.

As a general rule, for prior art to anticipate under section 102, every element of the claimed invention must be identically disclosed in a single reference. Corning Glass Works v. Sumitomo Electric, 9 U.S.P.Q.2d 1962, 1965 (Fed. Cir. 1989). The exclusion of a claimed element, no matter how insubstantial or obvious, from a reference is enough to negate anticipation. Connell v. Sears, Roebuck & Co., 220 U.S.P.Q 193, 1098 (Fed. Cir. 1983). Applicants respectfully submit that Yguerabide cannot be applied to support an anticipation rejection of the new claims under 35 U.S.C. section 102(e).

Specifically, the Examiner alleged that Yguerabide taught detection and measurement of one or more analytes in a sample using particles of specific composition and size using light scattering. The discussion is found starting in col. 82, line 35, of Yguerabide. Col. 83 provides further discussion regarding particle size and particle binding to a surface. There is no discussion of any aggregate probe or method of use thereof. Aggregate probes including two or more types of nanoparticles have been found to be surprisingly useful in target nucleic acid detection assays because they eliminate the need for buildup of multiple layers of individual nanopaticles in order to obtain or enhance a detectable change resulting from the binding of a nanoparticle oligonucleotide conjugate to a target nucleic acid. See the specification at page 49, line 6-page 50, line 2. Furthermore, new claims 433-465 recites limitations that are neither taught, made obvious, or suggested by the cited reference. Thus, the Applicant respectfully submits that Yguerabide cannot be applied to support a section 102(e) rejection of the new claims.

In conclusion, the Applicants respectfully submit that the claims in this application are in allowable condition and request a Notice to this effect.

Reconsideration of this application is respectfully requested and a favorable determination is earnestly solicited. The Examiner is invited to contact the undersigned

representative if the Examiner believes that this would be helpful in expediting the prosecution of this application.

Respectfully submitted,

**Emily Miao** 

Reg. No. 35,285

Dated: <u>001, 3, 2003</u>

McDonnell Boehnen Hulbert & Berghoff, Ltd. 300 South Wacker Drive Chicago, IL 60606

Telephone: 312-913-0001 Facsimile: 312-913-0001

## **APPENDIX A**

ATTY	Serial No./		
Case No.	Filing Date	Inventors/Title	Status
00-653-A	U.S. 09/927,777	Mirkin, Letsinger,	PENDING
	Filed 8/10/01	Mucic, Storhoff,	
		Elghanian, Taton,	
		Garamella, Li, Park/	
		NANOPARTICLES	
		HAVING	
		OLIGONUCLEOTI	
		DES ATTACHED	
		THERETO AND	
		USES THEREFORE	
00-713-B1	09/923,625	Mirkin, Letsinger,	PENDING
	Filed 8/7/01	Mucic, Storhoff,	
		Elghanian/	
		NANOPARTICLES	
		HAVING	
		OLIGONUCLEOTI	
		DES ATTACHED	
		THERETO AND	
		USES THEREFOR	-
00-713-C	09/344,667, filed	Mirkin, Letsinger,	U.S. Patent No.
	6/25/99	Mucic, Storhoff,	6,361,944, issued
		Elghanian/	3/26/02
		NANOPARTICLES	
		HAVING	
		OLIGONUCLEOTI	
		DES ATTACHED	
		THERETO AND USES THEREFORE	
00-713-I	U.S.S.N	Mirkin, Letsinger,	U.S. Patent No.
00-/13-1	09/603,830	Mucic, Storhoff,	6,506,564, issued
	Filed 6/26/00	Elghanian, Taton;	1/14/03
	1 11cd 0/20/00	NANOPARTICLES	1/14/05
		HAVING	
		OLIGONUCLEOTI	
		DES ATTACHED	
		THERETO AND	
		USES THEREFOR	
00-713-I-1	09/961,949	Mirkin, Letsinger,	U.S. Patent No.
<del>-</del>	9/20/01	Mucic, Storhoff,	6,582,921, issued
		Elghanian, Taton;	June 24, 2003

ATTY	Serial No./		
Case No.	Filing Date	Inventors/Title	Status
		NANOPARTICLES	
		HAVING	
		OLIGONUCLEOTI	
		DES ATTACHED	
		THERETO AND	
	00/055 010	USES THEREFOR	DEN ID D IC
00-713-I-2	09/957,318 9/20/01	See 00-713-I-1	PENDING
00-713-I-3	09/957,313 9/20/01	See 00-713-I-1	ALLOWED
00-713-I-4	09/966,491	See 00-713-I-1	U.S. Patent No.
	9/28/01		6,610,491
00-713-I-5	09/966,312 9/28/01	See 00-713-I-1	ALLOWED
00-713-I-6	09/967,409 9/28/01	See 00-713-I-1	PENDING
00-713-I-7	09/974,500 10/10/01	See 00-713-I-1	PENDING
00-713-I-8	09/974,007 10/10/01	See 00-713-I-1	PENDING
00-713-I-9	09/973,638 10/10/01	See 00-713-I-1	PENDING
00-713-I-	09/973,788	See 00-713-I-1	ALLOWED
10	10/10/01		
00-713-I- 11	09/975,062 10/11/01	See 00-713-I-1	ALLOWED
00-713-I- 12	09/975,376 10/11/01	See 00-713-I-1	PENDING
00-713-I- 13	09/975,384 10/11/01	See 00-713-I-1	PENDING
00-713-I- 14	09/975,498 10/11/01	See 00-713-I-1	ALLOWED

Case No.         Filing Date         Inventors/Title         Status           00-713-I- 15         09/975,059 11/11/01         See 00-713-I-1         PENDING           00-713-I- 16         09/976,601 10/12/01         See 00-713-I-1         PENDING           00-713-I- 17         09/976,968 10/12/01         See 00-713-I-1         PENDING           00-713-I- 18         09/976,971 10/12/01         See 00-713-I-1         ALLOWED           00-713-I- 19         09/976,863 10/12/01         See 00-713-I-1         PENDING	ATTY	Serial No./		T ag
15       11/11/01       See 00-713-I-1       PENDING         00-713-I- 16       10/12/01       See 00-713-I-1       PENDING         00-713-I- 09/976,968 17       See 00-713-I-1       PENDING         00-713-I- 09/976,971 18       See 00-713-I-1       ALLOWED         00-713-I- 09/976,863       See 00-713-I-1       PENDING	Case No.	Filing Date	Inventors/Title	Status
15       11/11/01       See 00-713-I-1       PENDING         00-713-I- 16       10/12/01       See 00-713-I-1       PENDING         00-713-I- 09/976,968 17       See 00-713-I-1       PENDING         00-713-I- 09/976,971 18       See 00-713-I-1       ALLOWED         00-713-I- 09/976,863       See 00-713-I-1       PENDING				
00-713-I- 16         09/976,601 10/12/01         See 00-713-I-1 See 00-713-I-1         PENDING           00-713-I- 17         09/976,968 10/12/01         See 00-713-I-1 See 00-713-I-1         PENDING           00-713-I- 18         09/976,971 10/12/01         See 00-713-I-1 See 00-713-I-1         ALLOWED           00-713-I- 10         09/976,863         See 00-713-I-1 See 00-713-I-1         PENDING			See 00-713-I-1	PENDING
16       10/12/01       See 00-713-I-1       PENDING         17       10/12/01       See 00-713-I-1       PENDING         00-713-I- 09/976,971 18       See 00-713-I-1       ALLOWED         00-713-I- 09/976,863       See 00-713-I-1       PENDING	15	11/11/01		
16       10/12/01       See 00-713-I-1       PENDING         17       10/12/01       See 00-713-I-1       PENDING         00-713-I- 09/976,971 18       See 00-713-I-1       ALLOWED         00-713-I- 09/976,863       See 00-713-I-1       PENDING	00 712 T	00/076 601	Sec. 00. 712 I.1	DENIDING
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17       10/12/01         00-713-I-       09/976,971       See 00-713-I-1       ALLOWED         18       10/12/01       See 00-713-I-1       PENDING	10	10/12/01		
00-713-I- 18         09/976,971 10/12/01         See 00-713-I-1 See 00-713-I-1         ALLOWED           00-713-I- 09/976,863         See 00-713-I-1 See 00-713-I-1         PENDING	00-713-I-	09/976,968	See 00-713-I-1	PENDING
18         10/12/01           00-713-I-         09/976,863           See 00-713-I-1         PENDING	17	10/12/01		
18         10/12/01           00-713-I-         09/976,863           See 00-713-I-1         PENDING				
<b>00-713-I-</b> 09/976,863 See 00-713-I-1 PENDING		· · · · · · · · · · · · · · · · · · ·	See 00-713-I-1	ALLOWED
	18	10/12/01		
	00-713-I-	09/976.863	See 00-713-I-1	PENDING
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<b>00-713-I-</b> 09/976,577 See 00-713-I-1 ALLOWED	00-713-I-	09/976,577	See 00-713-I-1	ALLOWED
20 10/12/01	20	10/12/01		
<b>00-713-I-</b> 09/976,618 See 00-713-I-1 PENDING			See 00-713-I-1	PENDING
21 10/12/01	21	10/12/01		
<b>00-713-I-</b> 09/981,344 See 00-713-I-1 PENDING	00-713-I-	09/981 344	See 00-713-I-1	PENDING
22 10/15/01 See 50 715 1 1 1 ENDING		,	500 00 715 1 1	LINDING
<b>00-713-I-</b> 09/976,900 See 00-713-I-1 PENDING	00-713-I-	09/976,900	See 00-713-I-1	PENDING
23 10/12/01	23	10/12/01		
00.710.1	00 510 7	00/076 617	0.00.712.11	DEMBRIC
00-713-I-         09/976,617         See 00-713-I-1         PENDING           24         10/12/01         Pending         Pending		-	See 00-713-1-1	PENDING
24   10/12/01	24	10/12/01		
<b>00-713-I-</b> 09/976,378 See 00-713-I-1 PENDING	00-713-I-	09/976,378	See 00-713-I-1	PENDING
25 10/12/01	25			
<b>00-713-i-</b> 10/410,324 See 00-713-I-1 PENDING			See 00-713-I-1	PENDING
26 04/10/03				110 2
00-713-L U.S.S.N. Mirkin, Letsinger, U.S. Patent No.	00-713-L			
09/693,005 Mucic, Storhoff, 6,495,324, issued		·		1
Filed 10/20/00   Elghanian/   12/17/02		rilea 10/20/00	1 0	12/1//02
NANOPARTICLES HAVING	•		i	
OLIGONUCLEOTI				
DES ATTACHED				
THERETO AND	•			

ATTY	Serial No./		
Case No.	Filing Date	Inventors/Title	Status
		USES THEREFORE	
00-713-M	U.S.S.N. 09/693,352 Filed 10/20/00	Mirkin, Letsinger, Mucic, Storhoff, Elghanian/ NANOPARTICLES HAVING OLIGONUCLEOTI DES ATTACHED THERETO AND USES THEREFORE	U.S. Patent No. 6,417,340, issued 7/9/02
00-714-G	U.S. 09/830,620 Filed 8/15/01	Mirkin, Nguyen/ NANOPARTICLES WITH POLYMER SHELLS	PENDING
00-715-A	U.S. 09/760,500 Filed 1/12/01	Mirkin, Letsinger, Mucic, Storhoff, Elghanian, Taton; Garamella, Li/ METHOD OF ATTACHING OLIGONUCLEOTI DES TO NANOPARTICLES AND PRODUCTS PRODUCED THEREBY	ALLOWED
00-1085-A	U.S.S.N. 09/820,279 Filed 3/28/01	Mirkin,Letsinger, etc./ METHOD AND MATERIALS FOR ASSAYING BIOLOGICAL MATERIALS	ALLOWED
00-1086-A	U.S. 09/903,461 Filed 7/11/01	Letsinger, Garimella/ METHOD OF DETECTION BY ENHANCEMENT OF SILVER STAINING	U.S. Patent No. 6,602,669, Filed 8/5/03
01-565-A	USSN 10/125,194 Filed 4/18/02	Mirkin, Nguygen, Watson, Park/ OLIGONUCLEOTI DE-MODIFIED ROMP POLYMERS AND CO-	PENDING

ATTY	Serial No./		
Case No.	Filing Date	Inventors/Title	Status
		POLYMERS	
01-599-A	U.S.S.N.	Storhoff/NOVEL	PENDING
	10/291,291	THIOL-BASED	
	Filed 11/08/02	METHOD FOR	
		ATTACHING	
		OLIGONUCLEOTI	
		DES TO	
		NANOPARTICLES	
01-661-A	U.S.S.N.	Mirkin, Cao, Jin/	PENDING
	10/034,451	DNA-MODIFIED	
	Filed 12/28/01	CORE-SHELL	
		AG/AU	
		NANOCRYSTALS	
01-661-C	U.S.S.N.	Mirkin, Cao, Jin/	PENDING
	10/153,483	DNA-MODIFIED	
	Filed 5/22/02	CORE-SHELL	
		AG/AU	
		NANOCRYSTALS	
01-661-E	U.S.S.N.	Mirkin, Cao, Jin/	PENDING
	10/397,579	DNA-MODIFIED	
	3/26/03	CORE-SHELL	
		AG/AU	
200.		NANOCRYSTALS	
01-1565-A	U.S.S.N.	Park, Taton,	PENDING
	10/266,983	Mirkin/ARRAY-	
	Filed 10/08/02	BASED	
		ELECTRICAL	
		DETECTION OF	
		DNA USING	
		NANOPARTICLE	
04.4=0=.	TICCOI	PROBES	DENIDDIC
01-1705-A	U.S.S.N.	Nam, Park,	PENDING
	10/108,211	Mirkin/BIO-	
	Filed 3/27/02	BARCODES BASED ON	
		OLIGONUCLEOTI	
		DE-MODIFIED	
		NANOPARTICLES	
02-338-B	USSN 10/172,428	Cao, Jin, Nam,	PENDING
U2-330-D	Filed 6/14/02	Mirkin/MULTICHA	
	1 1100 0/14/02	NNEL DETECTION	
		USING	
		NANOPARTICLE	
		PROBES WITH	
		I RODES WITH	<u> </u>

ATTY	Serial No./		
Case No.	Filing Date	Inventors/Title	Status
		RAMAN	
		SPECTROSCOPIC	
		FINGERPRINTS	
02-338-C	10/431,341	Cao, Jin, Nam,	PENDING
	5/7/03	Mirkin/MULTICHA	
		NNEL DETECTION	
		USING	
		NANOPARTICLE	
		PROBES WITH	
		RAMAN	
		SPECTROSCOPIC	
		FINGERPRINTS	